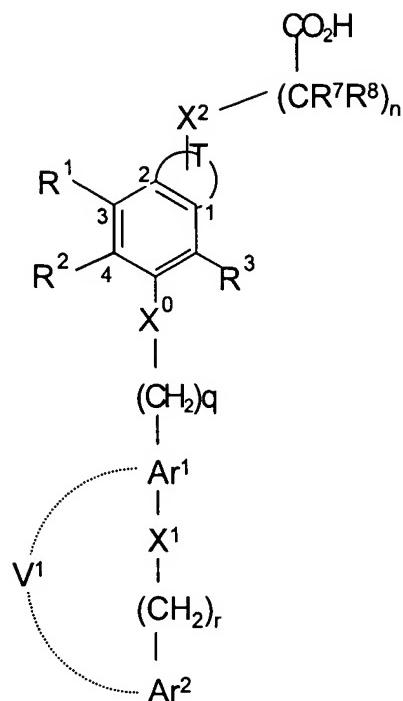


What is claimed is:

1. A compound having a Formula (I),



5

Formula I

a pharmaceutically acceptable salt, ester, amide or prodrug thereof, or a pharmaceutically
10 acceptable salt of the prodrug wherein:

X^0 and X^1 are each independently absent, O, S, NR⁴, -CH₂-CH₂-, -CH=CH, or
 $\text{---C}\equiv\text{C---}$;

Ar^1 and Ar^2 are each independently absent or unsubstituted or substituted aryl or
heteroaryl,

is absent; or when present, is a saturated or unsaturated hydrocarbon chain which is substituted or unsubstituted, wherein said chain has from 1 to 4 atoms so

that , Ar¹, X¹, (CH₂)_r and Ar², together form a five to eight membered ring;

- 5 T is a saturated or unsaturated, substituted or unsubstituted hydrocarbon chain or hydrocarbon-heteroatom chain having from 3 to 6 atoms wherein the carbon atom of position 1 is connected to the carbon atom of position 2 to form a five to eight member

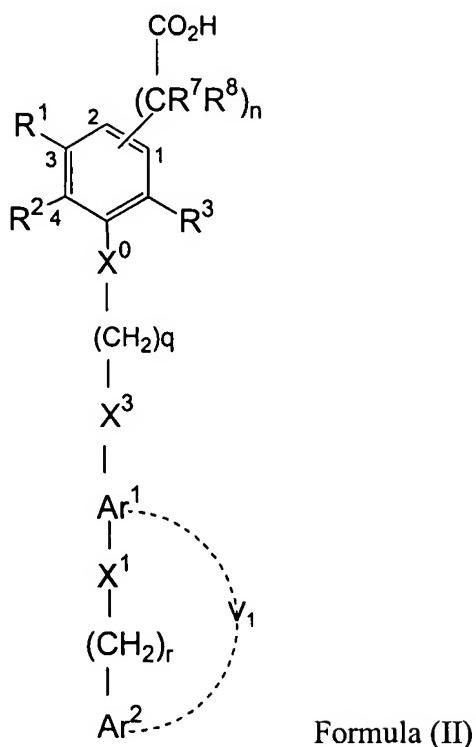
$$\begin{array}{c} \text{CO}_2\text{H} \\ | \\ \text{ring wherein the } \text{X}^2 - (\text{CR}^7\text{R}^8)_n \text{ is attached to a substitutionally available position of said ring;} \end{array}$$

- X² is absent, O, S, or NR⁴;
- 10 R¹, R², and R³ are independently hydrogen, lower alkyl, lower alkoxy, lower thioalkoxy, -O(CH₂)_pCF₃, halogen, nitro, cyano, -OH, -SH, -CF₃, S(O)_pAlkyl, S(O)_pAryl, -(CH₂)_mOR⁴, or -(CH₂)_mNR⁵R⁶, COR⁴, -CONR⁵R⁶, -CO₂R⁴, or -NR⁵R⁶ or R¹ and R² are joined together to form a substituted or unsubstituted, saturated or unsaturated cyloalkyl or heterocycloalkyl ring;

- 15 R⁴ is hydrogen, alkyl, alkenyl, alkynyl, acyl, SO₂Aryl, SO₂Alkyl or aryl; R⁵ and R⁶ are independently hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, SO₂Alkyl aryl or SO₂Aryl, or joined together to form a 4 to 7 member ring having 0 to 3 heteroatoms;

- R⁷ and R⁸ are independently H, lower alkyl, halo, or R⁷ and R⁸ taken together form a 3-6 membered hydrocarbon ring, optionally containing a heteroatom;
- m is an integer from 0 to 5;
- n is an integer from 0 to 5;
- p is an integer from 0 to 2.
- q is an integer from 0 to 10; and
- 25 r is an integer from 0 to 10.

2. A compound having a Formula II,



5 a pharmaceutically acceptable salt, ester, amide or prodrug thereof or a pharmaceutically acceptable salt of the prodrug wherein:

X^3 is O, C=O, S, CHOR¹¹ where R¹¹ is lower alkyl, aryl, acyl, -SO₂alkyl- or -SO₂aryl, absent or NR⁴; R¹, R², and R³ are independently hydrogen, lower alkyl, lower alkoxy, lower thioalkoxy, -O(CH₂)_pCF₃, halogen, nitro, cyano, -OH, -SH, -CF₃,

10 S(O)_pAlkyl, S(O)_pAryl, -(CH₂)_mOR⁴, or -(CH₂)_mNR⁵R⁶, COR⁴, -CONR⁵R⁶, -CO₂R⁴, or -NR⁵R⁶ or R¹ and R² are joined together to form a substituted or unsubstituted, saturated or unsaturated cycloalkyl or heterocycloalkyl ring;

R⁴ is hydrogen, alkyl, alkenyl, alkynyl, acyl, SO₂Aryl, SO₂Alkyl or aryl;

R⁵ and R⁶ are independently hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl,

15 SO₂Alkyl aryl or SO₂Aryl, or joined together to form a 4 to 7 member ring having 0 to 3 heteroatoms;

R⁷ and R⁸ are independently H, lower alkyl, halo, or R⁷ and R⁸ taken together form a 3-6 membered hydrocarbon ring, optionally containing a heteroatom;

X^0 and X^1 are each independently absent, O, S, NR^4 , $-CH_2-CH_2-$, $-CH=CH$, or
 $-C\equiv C-$; X^2 is absent, O, S, or NR^4 ;

Ar^1 and Ar^2 are each independently absent or unsubstituted or substituted aryl or

5 heteroaryl,

is absent; or when present, is a saturated or unsaturated hydrocarbon chain which is substituted or unsubstituted, wherein said chain has from 1 to 4 atoms so

that , Ar^1 , X^1 , $(CH_2)_r$, and Ar^2 , together form a five to eight membered ring;
n is an integer from 0 to 5; q is an integer from 0 to 10; and r is an integer from 0 to 10.

10

3. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

X^0 is S or O;

X^1 is absent, O or S;

15 Ar^1 and Ar^2 are each independently absent, or unsubstituted or substituted aryl or heteroaryl;

is absent;

q is 1; and

r is 0 or 1.

20

4. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein: X^0 is S or O; X^1 is O or absent; and Ar^1 and Ar^2 are each independently unsubstituted or substituted aryl or heteroaryl.

5. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

T is $-\text{CH}_2\text{CH}_2\text{CO}-$, $-\text{CH}_2\text{-CH}_2\text{-O-CO-}$, $-\text{CH}_2\text{-CH}_2\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{HC=CH-}$
5 HC=CH- , $-\text{N=CH-HC=CH-}$, $-\text{HC=N-HC=CH-}$, $-\text{HC=CH-N=CH-}$, $-\text{HC=CH-HC=N-}$, $-\text{CH}_2\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-O-CH}_2\text{-}$, $-\text{CH}_2\text{-HC=CH-}$, $-\text{CH}_2\text{CH}_2\text{-NH-CH}_2\text{-}$, $-\text{COCH=CH-}$
 O- , $-\text{O-CH=CH-CO-}$, $-\text{O-CH=CH-}$, $-\text{CH=CH-O-}$, $-\text{O-CH}_2\text{-CH=CH-}$, $-\text{CH=CH-CH}_2\text{-O-}$, $-\text{CH}_2\text{-CH}_2\text{-CO-NR}^4$, $-\text{CH}_2\text{-CH}_2\text{-CO-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-NR}^4\text{-CH}_2\text{-}$, $-\text{CH}_2\text{-NR}^4\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{CH=CH-NR}^4\text{-}$, $-\text{NR}^4\text{-CH=CH-}$, $-\text{CH=CH-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-NR}^4\text{-}$, $-\text{NR}^4\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{O-CH}_2\text{-CH}_2\text{-}$,
10 $-\text{O-CH}_2\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-O-}$, $-\text{CH}_2\text{-CH}_2\text{-CH}_2\text{-O-}$, $-\text{O-CH(CH}_3\text{)-CH}_2\text{-}$
 $\text{CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-CH(CH}_3\text{)-O-}$, $-\text{CH}_2\text{-CH}_2\text{-CH}_2\text{-NR}^4\text{-}$, $-\text{NR}^4\text{-CH}_2\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-CO-NR}^4\text{-}$, $-\text{NR}^4\text{-CO-CH}_2\text{-CH}_2\text{-}$, $-\text{O- NR}^4\text{-CO-}$, $-\text{CO- NR}^4\text{-O-}$, $-\text{O-CH}_2\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-O-}$, $-\text{CH}_2\text{-CH}_2\text{-NR}^4\text{-CO-}$, $-\text{CH}_2\text{-CH}_2\text{-CH}_2\text{-CO-}$, $-\text{CO-CH}_2\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{NR}^4\text{-CO-CH}_2\text{-CH}_2\text{-}$, $-\text{CO-NR}^4\text{-CH}_2\text{-CH}_2\text{-}$, $-\text{CH}_2\text{-CH}_2\text{-CO-}$, $-\text{CH}_2\text{-CO-CH}_2\text{-}$, $-\text{CO-CH}_2\text{-CH}_2\text{-S-}$
15 C=C- , $-\text{C=C-S-}$, $-\text{S-C-C-}$, $-\text{C-C-S-}$,
 $-\text{S-C-C-C-}$, $-\text{C-C-C-S-}$, $-\text{C=C-C-S-}$, $-\text{C-C=C-S-}$, $-\text{S-C=C-C-}$, or $-\text{S-C-C=C-}$.

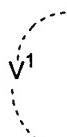
6. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein :

20 X^0 is S;

X^1 is absent;

Ar^1 is substituted phenyl;

Ar^2 is phenyl;


 V^1

is absent;

25 q is 1; and

r is 0 or 1.

7. A compound of claim 3, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

T is substituted with 1 or more substituents selected from the group consisting of lower alkyl, lower alkoxy, lower thioalkoxy, -O(CH₂)₀₋₂CF₃, halogen, nitro, cyano, =O, =S, -OH, -SH, -CF₃, -CO₂H, -CO₂C₁-C₆ alkyl, -NH₂, -NHC₁-C₆ alkyl, -CONR'R'', or -N(C₁-C₆alkyl)₂; and

5 R' and R'' are independently alkyl, alkenyl, alkynyl, aryl, or joined together to form a 4 to 7 member ring.

8. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein q is 1.

10

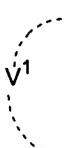
9. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein Ar¹ is substituted or unsubstituted phenyl.

15

10. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein Ar² is 4-trifluoromethylphenyl.

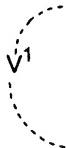
20

11. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or

prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein  is absent.

12. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or

25

prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein  is (CH₂)_t and t is an integer from 1 to 4.

13. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein

v1

is substituted with at least one substituent selected from the group consisting of lower alkyl, lower alkoxy, lower thioalkoxy, -O(CH₂)₀₋₂CF₃, halogen, nitro, cyano, =O, =S, -OH, -SH, -CF₃, -OCF₃, -CO₂H, -CO₂C_{1-C₆} alkyl, -NH₂, -NHC_{1-C₆} alkyl, -CONR'R'', or -N(C_{1-C₆} alkyl)₂ where R' and R'' are independently alkyl, akenyl, alkynyl, aryl, or joined together to form a 4 to 7 member ring.

10 14. A pharmaceutical composition comprising a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; and a pharmaceutically acceptable carrier, diluent, or vehicle.

15 15. A method of treating, preventing or controlling non-insulin dependent diabetes mellitus in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

20 25 16. A method of treating, preventing or controlling obesity in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

17. A method of treating, preventing or controlling hyperglycemia in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of

5 Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

18. A method of treating, preventing or controlling hyperlipidemia in a mammal comprising administering to the mammal in need thereof a therapeutically effective

10 amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

15 19. A method of treating, preventing or controlling hypercholesterolemia in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

20. A method of treating, preventing or controlling atherosclerosis in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of

25 Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

30 21. A method of treating, preventing or controlling hypertriglyceridemia in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or

prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

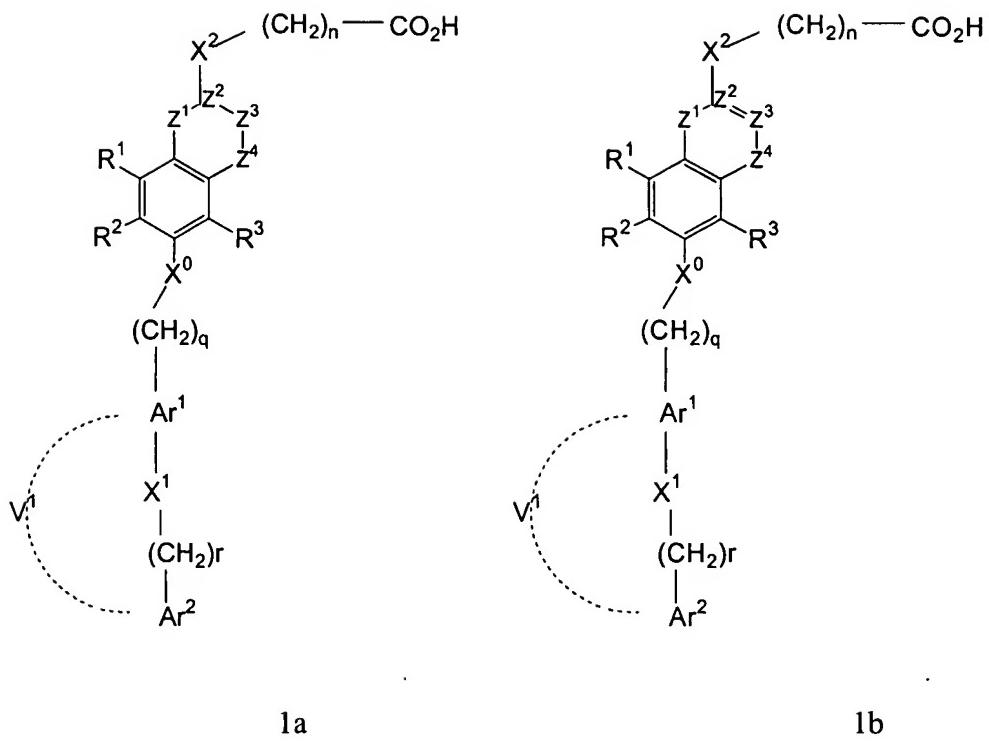
5 22. A method of treating, preventing or controlling hyperinsulinemia in a mammal comprising administering to the mammal in need thereof a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the
10 pharmaceutically acceptable salt of the prodrug.

15 23. A method of treating a patient exhibiting glucose disorders associated with circulating glucocorticoids, growth hormone, catecholamines, glucagon, or parathyroid hormone, comprising administering to the patient a therapeutically effective amount of a compound of Claim 1, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug; or a compound of Claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug.

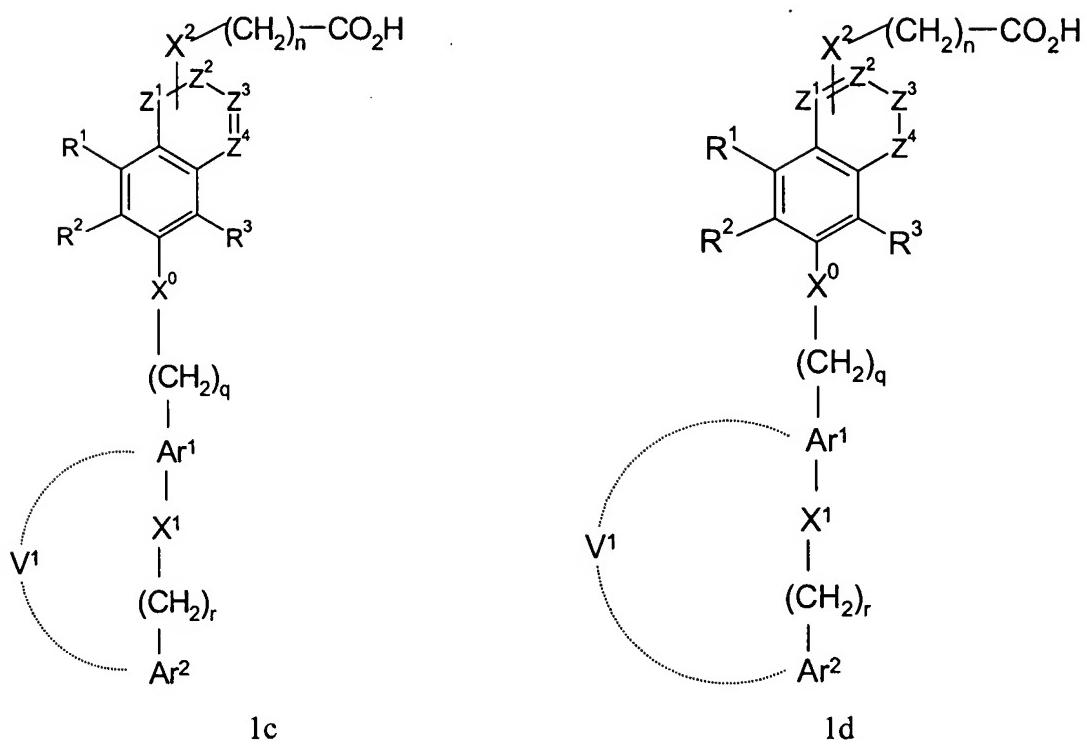
20 24. A compound of claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein X^3 is NR^4 or $C=O$.

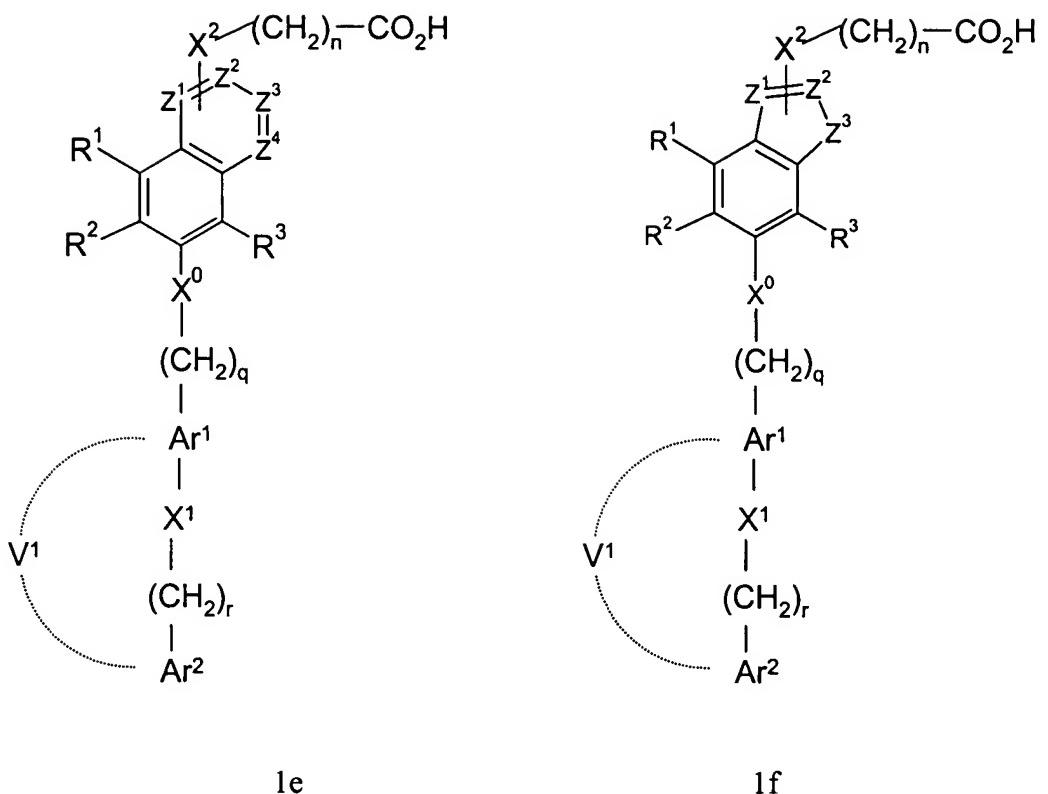
25 25. A compound of claim 2, the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein Ar^2 is chloro-phenyl, dichloro-phenyl-, trichlorophenyl, fluoro-phenyl-, difluorophenyl, trifluorophenyl, trifluoromethyl-phenyl, or fluoro-trifluoromethyl-phenyl-; and wherein Ar^1 is absent.

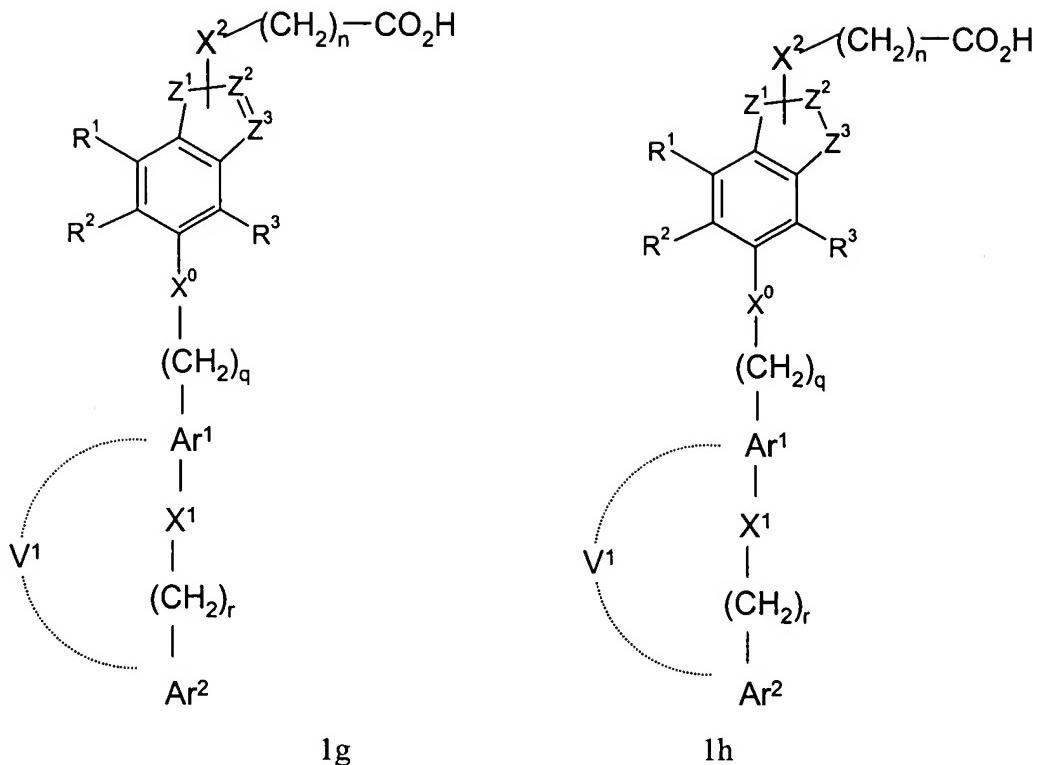
30 26. A compound of claim 1 having Formula 1a, Formula 1b, Formula 1c, Formula 1d, Formula 1e, Formula 1f, Formula 1g, or Formula 1h,



5







5

or the pharmaceutically acceptable salt, ester, amide or prodrug thereof, or the pharmaceutically acceptable salt of the prodrug wherein:

- 10 X⁰ is O or S;
- X² is absent, O, S, or NR⁴;
- R¹, R², and R³ are independently hydrogen, lower alkyl, lower alkoxy, lower thioalkoxy, -O(CH₂)_pCF₃, halogen, nitro, cyano, -OH, -SH, -CF₃, S(O)_pAlkyl, S(O)_pAryl, -(CH₂)_mOR⁴, or -(CH₂)_mNR⁵R⁶, COR⁴, -CO₂H, -CO₂R⁴, or -NR⁵R⁶ or R¹ and R² are joined together to form a substituted or unsubstituted, saturated or unsaturated cycloalkyl or heterocycloalkyl ring;

- 15 R⁴ is hydrogen, alkyl, alkenyl, alkynyl, or aryl;
- R⁵ and R⁶ are independently hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, SO₂Alkyl or, SO₂Aryl, or joined together to form a 4 to 7 member ring having 0 to 3 heteroatoms;
- m is an integer from 0 to 5;

n is an integer from 0 to 5;
p is an integer from 0 to 2,
 Z^1, Z^2, Z^3 , and Z^4 are independently O, S, CR^5R^6 , NR^{11} , or N;
 R^{11} is lower alkyl, acyl, aralkyl, $-SO_2$ alkyl, or $-SO_2Ar$, and wherein
5 Z^1, Z^2, Z^3 , and Z^4 are bonded to a sufficient number of hydrogen atoms or
substituents to complete the valency of each atom with the proviso that Z^1, Z^2, Z^3 , and Z^4
are not all heteroatoms and that not more than two adjacent atoms in Z^1, Z^2, Z^3 , and Z^4
are heteroatoms and that in Formulae 1b, 1c, 1d, 1f, and 1g, Z^1, Z^2, Z^3 , and Z^4 are not all
carbon atoms; and $X^1, Ar^1, Ar^2, \dots, r$ and q are as defined in claim 1.

10

27. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or
prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

R^1, R^2 , and R^3 are independently hydrogen, alkyl, or alkoxy.

15 28. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or
prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

R^1 and R^3 are hydrogen; and

R^2 is alkyl or alkoxy.

20 29. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or
prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

R^1 and R^3 are hydrogen; and

R^2 is alkoxy.

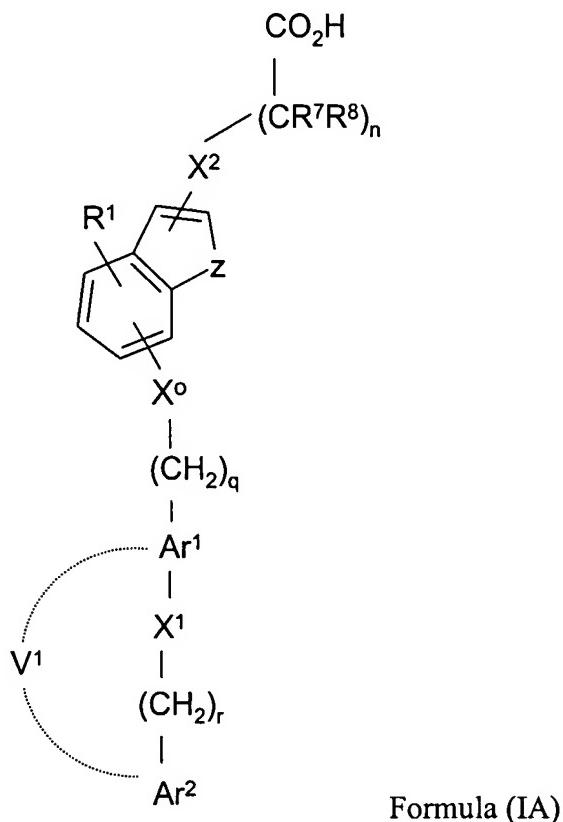
25 30. A compound of claim 1, the pharmaceutically acceptable salt, ester, amide or
prodrug thereof, or the pharmaceutically acceptable salt of the prodrug, wherein:

R^1 and R^3 are independently hydrogen, methyl, ethyl, isopropyl, n-propyl, t-butyl,
n-butyl, or isobutyl; and

R^2 is methyoxy, ethoxy, isopropoxy, n-propoxy, t-butoxy, n-butoxy, or isobutoxy.

31. A compound selected from the group consisting of: {6-(4'-Trifluoromethyl-biphenyl-4-ylmethylsulfanyl)-chroman-2-yl}-acetic acid; {6-[4-(5-Trifluoromethyl-pyridin-2-yl)-benzylsulfanyl]-chroman-2-yl}-acetic acid; {6-[4-(2,5-Dichloro-
5
benzyloxy)-benzylsulfanyl]-chroman-2-yl}-acetic acid; {6-[4-(4-Trifluoromethyl-benzyloxy)-benzylsulfanyl]-chroman-2-yl}-acetic acid; {6-[5-(4-Trifluoromethyl-phenyl)-isoxazol-3-ylmethylsulfany]-chroman-2-yl}-acetic acid; {6-[3-(4-Trifluoromethyl-benzyloxy)-benzylsulfanyl]-chroman-2-yl}-acetic acid; {6-[2-(4-Trifluoromethyl-phenyl)-thiazol-4-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; {6-[4-(4-Trifluoromethyl-benzyloxy)-benzyloxy]-benzo[b]thiophen-3-yl}-acetic acid; {6-[2-(4-Trifluoromethyl-benzyloxy)-benzyloxy]-benzo[b]thiophen-3-yl}-acetic acid; and
10
pharmaceutically acceptable salts thereof.

32. A compound having a formula (IA),



a pharmaceutically acceptable salt, ester, amide, or prodrug thereof, or a pharmaceutically acceptable salt of the prodrug wherein:

Z=S, O or NR⁴, Ar¹ and Ar² are each independently absent or unsubstituted aryl or heteroaryl,

5 is absent; or when present, is a saturated or unsaturated hydrocarbon chain which is substituted or unsubstituted, wherein said chain has from 1 to 4 atoms so

that , Ar¹, X¹, (CH₂)_r and Ar², together form a five to eight membered ring; X⁰ and X¹ are each independently absent, O, S, NR⁴, -CH₂-CH₂-, -CH=CH, or

—C≡C—; X² is absent, O, S, or NR⁴; R¹ is independently hydrogen, lower alkyl, lower
10 alkoxy, lower thioalkoxy, -O(CH₂)_pCF₃, halogen, nitro, cyano, -OH, -SH, -CF₃,
S(O)_pAlkyl, S(O)_pAryl, -(CH₂)_mOR⁴, -(CH₂)_mNR⁵R⁶, COR⁴, -CONR⁵R⁶, -CO₂R⁴, or
-NR⁵R⁶;

R⁴ is hydrogen, alkyl, alkenyl, alkynyl, acyl, SO₂Aryl, SO₂Alkyl or aryl;

R⁵ and R⁶ are independently hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl, SO₂Alkyl aryl

15 or SO₂Aryl, or joined together to form a 4 to 7 member ring having 0 to 3 heteroatoms; R⁷ and R⁸ are independently H, lower alkyl, halo, or R⁷ and R⁸ taken together form a 3-6 membered hydrocarbon ring, optionally containing a heteroatom;

n is an integer from 0 to 5;

q is an integer from 0 to 10; and

20 r is an integer from 0 to 10.

33. A compound selected from the group consisting of: {6-[4-(4-Trifluoromethyl-benzyloxy)-benzyloxy]-benzo[b]thiophen-3-yl}-acetic acid; {6-[5-(4-Trifluoromethyl-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; {6-[3-(4-

25 Trifluoromethyl-benzyloxy)-benzyloxy]-benzo[b]thiophen-3-yl}-acetic acid; {6-[2-(4-Trifluoromethyl-phenyl)-thiazol-4-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; {6-

[5-(4-Chloro-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; and pharmaceutically acceptable salts thereof.

34. A compound of claim 32, the pharmaceutically acceptable amide ester or prodrug
5 thereof, or the pharmaceutically acceptable salt of the prodrug wherein:

X^0 is oxygen;

X^1 is absent or O;

Ar^1 is a substituted or unsubstituted aryl or heteroaryl;

Ar^2 is a substituted phenyl;

V¹

is absent;

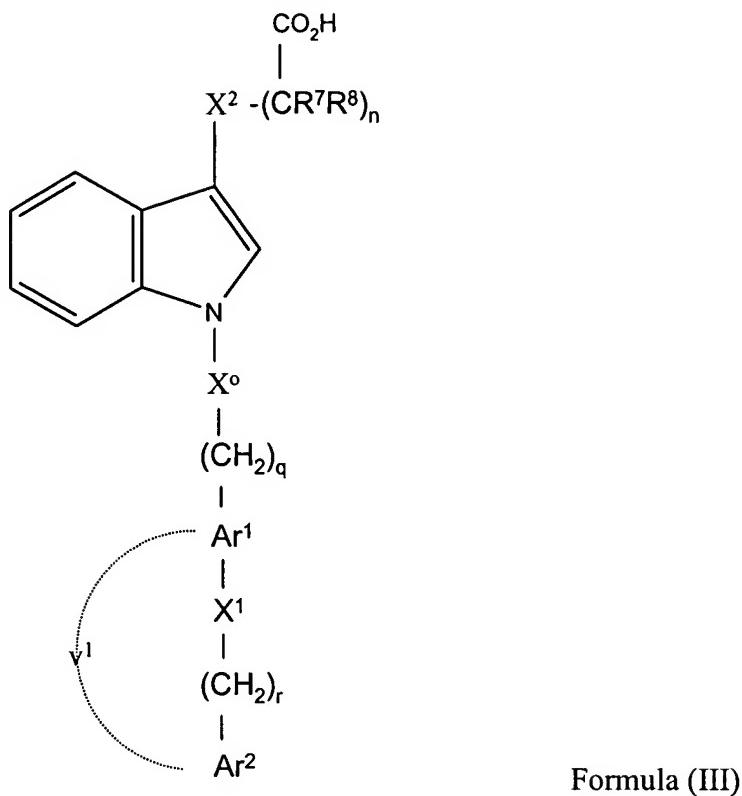
X^2 is absent

n is an integer from 0 to 5

q is an integer from 0 to 3; and

r is an integer from 0 to 3.

35. A compound having a Formula III,



5 a pharmaceutically salt, ester amide or prodrug thereof, or a pharmaceutically acceptable salt of the prodrug wherein:

Ar^1 and Ar^2 are each independently absent or unsubstituted or substituted aryl or heteroaryl,

10 V^1 is absent; or when present, V^1 is a saturated or unsaturated hydrocarbon chain which is substituted or unsubstituted, wherein said chain has from 1 to 4 atoms so

that V^1 , Ar^1 , X^1 , $(\text{CH}_2)_r$ and Ar^2 , together form a five to eight membered ring;

X^0 and X^1 are each independently absent, O, S, NR^4 , $-CH_2-CH_2-$, $-CH=CH$, or $-C\equiv C-$; X^2 is absent, O, S, or NR^4 ;

R^4 is hydrogen alkyl, alkenyl, alkynyl, acyl, SO_2Ar , SO_2Alkyl or aryl;

R^7 and R^8 are independently H, lower alkyl, halo, or R^7 and R^8 taken together form a 3-6

5 membered hydrocarbon ring, optionally containing a heteroatom;

n is an integer from 0 to 5;

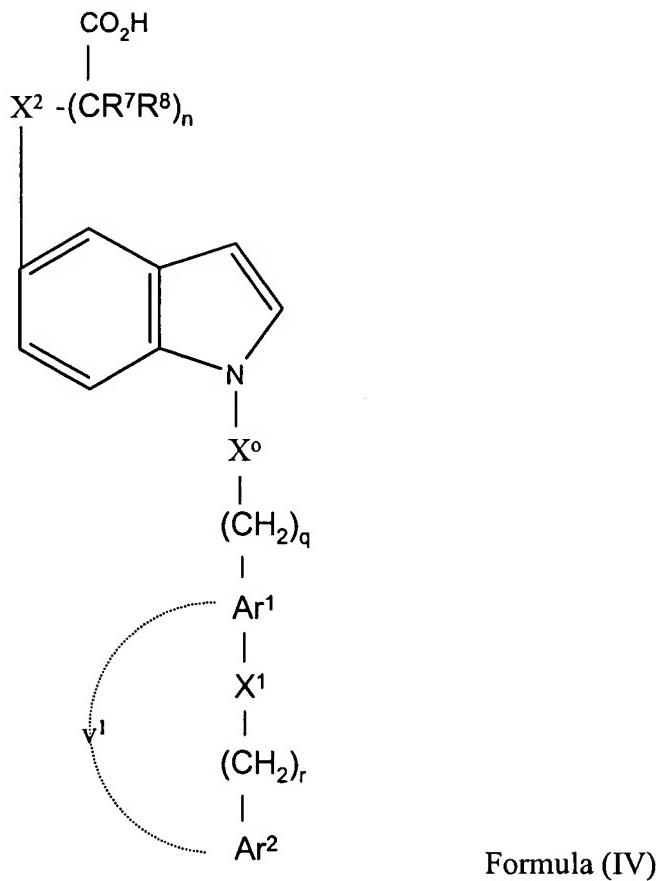
q is an integer from 0 to 10; and

r is an integer from 0 to 10.

- 10 36. A compound selected from the group consisting of: 3-[1-[3-(4-Trifluoromethyl-benzyloxy)-benzyl]-1H-indol-3-yl]-propionic acid; 3-[1-[4-(4-Trifluoromethyl-benzyloxy)-benzyl]-1H-indol-3-yl]-propionic acid; 3-[1-(4'-Trifluoromethyl-biphenyl-4-ylmethyl)-1H-indol-3-yl]-propionic acid; {1-[3-(4-Trifluoromethyl-benzyloxy)-benzyl]-1H-indol-3-yl}-acetic acid, and pharmaceutically acceptable salts thereof.

15

37. A compound having a Formula IV,



- 5 a pharmaceutically acceptable salt, ester, amide, or prodrug thereof, or a pharmaceutically acceptable salt of the prodrug wherein:
 Ar^1 and Ar^2 are each independently absent or unsubstituted or substituted aryl or heteroaryl,
V¹ is absent; or when present, V¹ is a saturated or unsaturated hydrocarbon
10 chain which is substituted or unsubstituted, wherein said chain has from 1 to 4 atoms so
V¹ that Ar^1 , X^1 , $(\text{CH}_2)_r$ and Ar^2 , together form a five to eight membered ring;

X^0 and X^1 are each independently absent, O, S, NR^4 , $-CH_2-CH_2-$, $-CH=CH$, or $-C\equiv C-$; X^2 is absent, O, S, or NR^4 ;

R^4 is hydrogen alkyl, alkenyl, alkynyl, acyl, SO_2Arlyl , SO_2Alkyl or aryl;

R^7 and R^8 are independently H, lower alkyl, halo, or R^7 and R^8 taken together form a 3-6

5 membered hydrocarbon ring, optionally containing a heteroatom;

n is an integer from 0 to 5;

q is an integer from 0 to 10; and

r is an integer from 0 to 10.

10 38. A compound selected from the group consisting of: [1-(4'-Trifluoromethyl-biphenyl-4-ylmethyl)-1H-indol-5-yloxy]-acetic acid, [1-(4'-Trifluoromethyl-biphenyl-4-ylmethyl)-1H-indol-4-yloxy]-acetic acid; and pharmaceutically acceptable salts thereof.

15 39. A compound selected from the group consisting of: [6-(4'-Trifluoromethyl-biphenyl-4-ylmethylsulfanyl)-chroman-2-yl]-acetic acid; {6-[5-(4-Trifluoromethyl-phenyl)-isoxazol-3-ylmethylsulfany]-chroman-2-yl}-acetic acid; {6-[2-(4-Trifluoromethyl-phenyl)-thiazol-4-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid; and pharmaceutically acceptable salts thereof.

20 40. A compound of claim 35, a pharmaceutically acceptable salt, amide, ester or prodrug thereof, or a pharmaceutically acceptable salt of the prodrug wherein:

X^0 is absent

X^1 is absent or O;

Ar^1 is a substituted or unsubstituted phenyl;

25 Ar^2 is 4-trifluoromethyl phenyl;

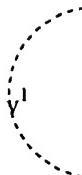


is absent;

X^2 is absent, 0, or S;
n is an integer from 0 to 5;
q is an integer from 0 to 3; and
r is an integer from 0 to 3.

5

41. A compound of claim 37, a pharmaceutically acceptable salt, amide, ester, prodrug thereof, or a pharmaceutically acceptable salt of the prodrug, wherein:
 X^0 and X^1 are absent;
 Ar^1 is a substituted or unsubstituted phenyl;
10 Ar^2 is 4-trifluoromethylphenyl;



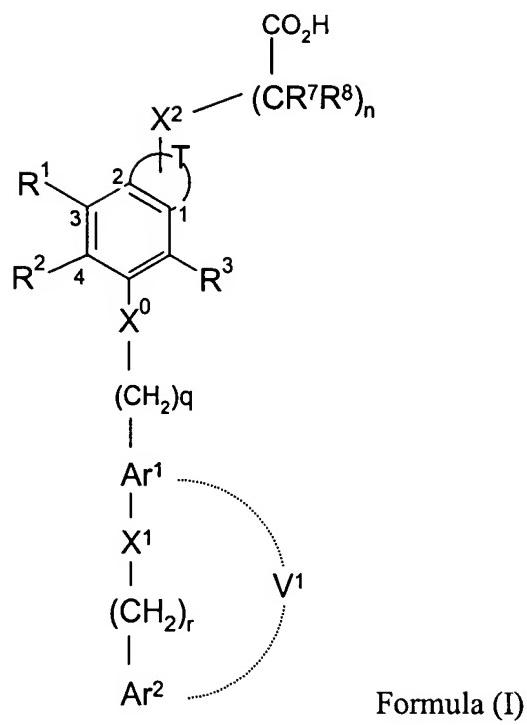
is absent;

X^2 is absent, 0 or S;
n is an integer from 0 to 5;
q is an integer from 0 to 3; and
15 r is an integer from 0 to 3.

42. A compound of claim 34, a pharmaceutically acceptable salt, ester, amide or prodrug thereof, or a pharmaceutically acceptable salt of the prodrug, wherein Ar^2 is trifluoromethyl-phenyl.

20

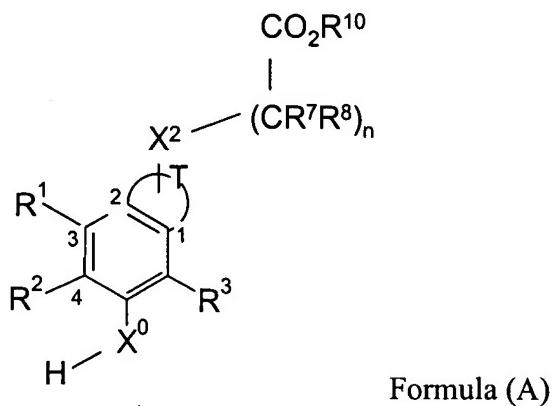
43. A method of making a compound of claim 1 having the Formula (I):



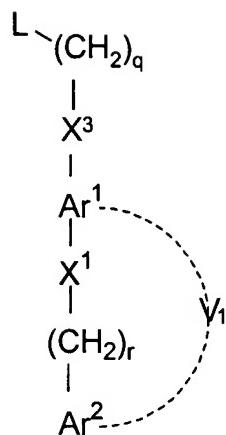
wherein X⁰, X¹, Ar¹, Ar², ..., T, X², R¹, R², R³, R⁴, R⁵, R⁶, R⁷, R⁸, m, n, p, q and r are as defined in claim 1.

comprising:

- 5 reacting a compound of Formula A,



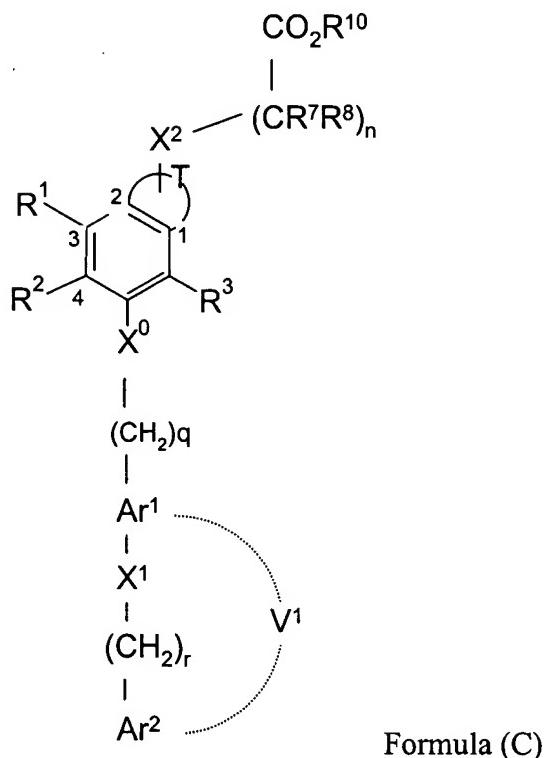
wherein R¹⁰ is a lower alkyl, with a compound of Formula B,



Formula (B)

wherein L is an appropriate leaving group and X^3 is absent, to form a compound having a Formula C,

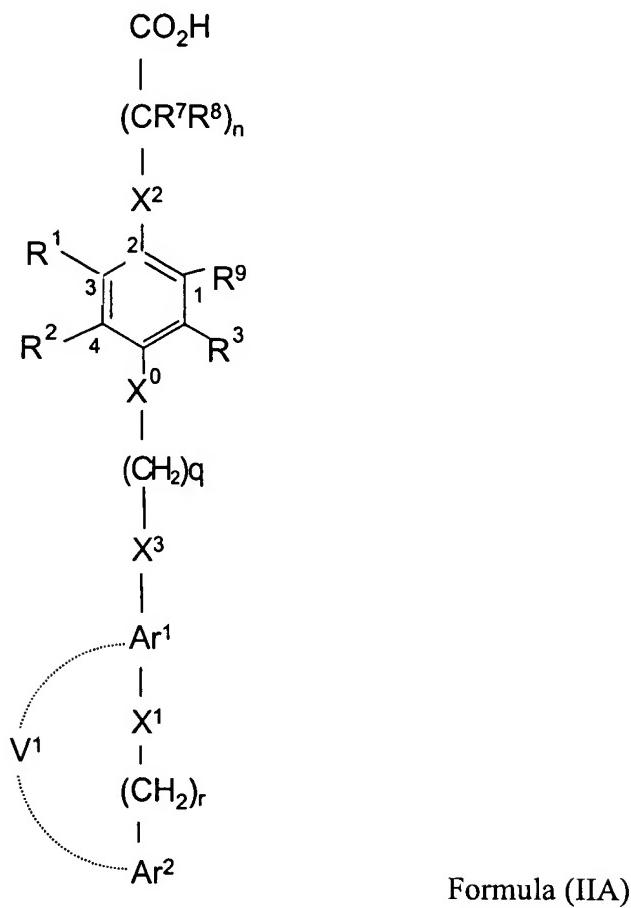
5



Formula (C)

and subsequently saponifying the compound having a Formula C to form the compound having the Formula I.

44. A compound having a Formula IIA,



5 a pharmaceutically acceptable salt, amide ester or prodrug thereof, or a pharmaceutically acceptable salt of the prodrug wherein:

X¹ and X³ are each independently O, C=O, S, CHOR¹¹, absent or NR⁴; R¹, R², R³ and R⁹ are independently hydrogen, lower alkyl, lower alkoxy, lower thioalkoxy, -O(CH₂)pCF₃, halogen, nitro, cyano, -OH, -SH, -CF₃, S(O)pAlkyl, S(O)pAryl, -(CH₂)mOR⁴,

10 -(CH₂)mNR⁵R⁶, -COR⁴, -CO₂H, -CO₂R⁴, or -NR⁵R⁶, or R¹ and R² are joined together to form a substituted or unsubstituted, saturated or unsaturated cycloalkyl or

heterocycloalkyl ring; R¹¹ is lower alkyl, aryl, acyl, -SO₂Alkyl, SO₂Aryl, absent or NR⁴;

X¹ and X⁰ are each independently absent, O, S, NR⁴, -CH₂-CH₂-, -CH=CH, or

—C≡C—; Ar¹ and Ar² are each independently absent or unsubstituted or substituted aryl

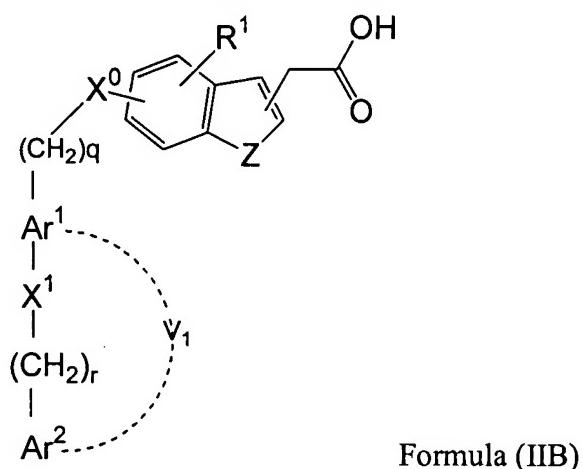
15 or heteroaryl,

X² is absent, O, S, or NR⁴;

R⁴ is hydrogen, alkyl, alkenyl, alkynyl, acyl, SO₂Aryl, SO₂Alkyl or aryl;
R⁵ and R⁶ are independently hydrogen, alkyl, alkenyl, alkynyl, cycloalkyl,
SO₂Alkyl aryl or SO₂Aryl, or joined together to form a 4 to 7 member ring having 0 to 3
heteroatoms;

- 5 R⁷ and R⁸ are independently H, lower alkyl, halo, or R⁷ and R⁸ taken together
form a 3-6 membered hydrocarbon ring, optionally containing a heteroatom;
m is an integer from 0 to 5;
n is an integer from 0 to 5;
p is an integer from 0 to 2.
10 q is an integer from 0 to 10; and
r is an integer from 0 to 10.

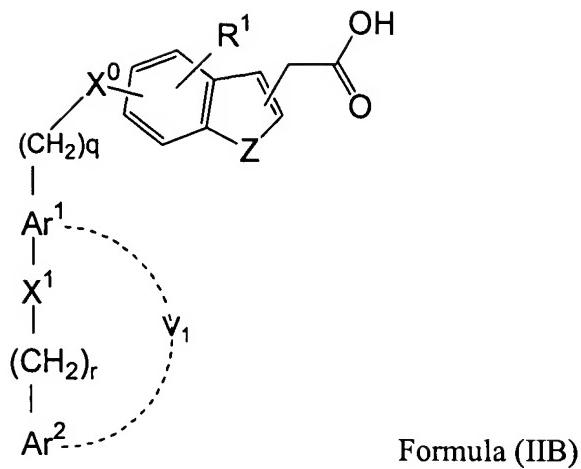
45. A compound of claim 32 having the Formula IIB,



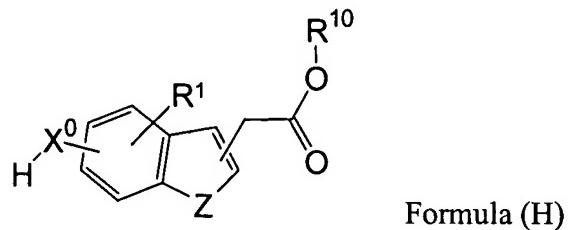
- 15 a pharmaceutically acceptable salt, ester, or amide thereof, or a pharmaceutically
acceptable salt of the prodrug wherein:

Z is NR⁴, S, or O; and R¹, R⁴, X⁰, X¹, Ar¹, Ar², ..., q and r are as defined in claim
32.

- 20 46. A method of making a compound of claim 45 having the Formula IIB

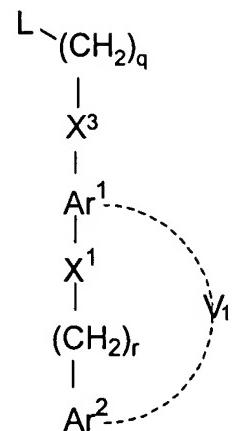


wherein Z , R^1 , R^4 , X^0 , X^1 , Ar^1 , Ar^2 , ..., q and r are as defined in claim 45,
comprising reacting a compound of Formula H wherein R^{10} is a lower alkyl,



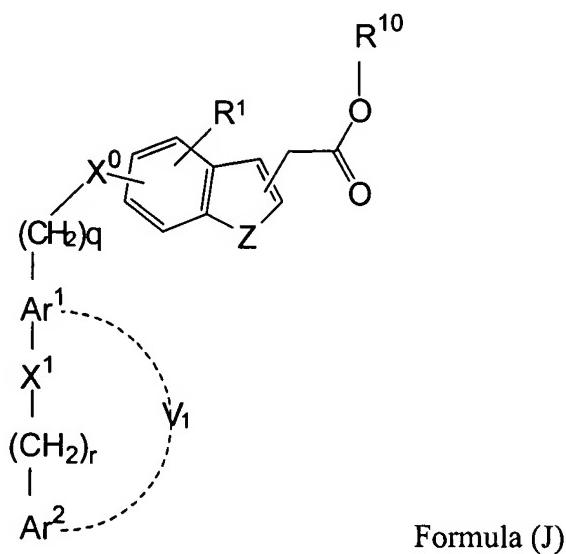
5

with a compound of Formula B:



$L = \text{leaving group}$ Formula (B)

wherein L is an appropriate leaving group, to form a compound of Formula J:



and subsequently saponifying the compound of Formula J to form the compound IIB.

- 5 47. A compound selected from the group consisting of :

{6-[5-(4-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;

{6-[5-(3-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;

10 {6-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;

{6-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;

{6-[5-(4-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzo[b]thiophen-3-

15 yl}-acetic acid;

{6-[5-(3-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzo[b]thiophen-3-yl}-acetic acid;

{6-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzo[b]thiophen-3-yl}-acetic acid;

20 {6-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzo[b]thiophen-3-yl}-acetic acid;

- {5-Methoxy-6-[5-(4-methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;
- {5-Methoxy-6-[5-(3-methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzo[b]thiophen-3-yl}-acetic acid;
- 5 {6-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-5-methoxy-benzo[b]thiophen-3-yl}-acetic acid;
- {6-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-5-methoxy-benzo[b]thiophen-3-yl}-acetic acid;
- {6-[5-(4-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzofuran-3-yl}-acetic acid;
- 10 {6-[5-(3-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzofuran-3-yl}-acetic acid;
- {6-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-benzofuran-3-yl}-acetic acid;
- {6-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-benzofuran-3-yl}-acetic acid;
- 15 {6-[5-(4-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzofuran-3-yl}-acetic acid;
- {6-[5-(3-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzofuran-3-yl}-acetic acid;
- {6-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzofuran-3-yl}-acetic acid;
- 20 {6-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-4-methyl-benzofuran-3-yl}-acetic acid;
- {5-Methoxy-6-[5-(4-methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzofuran-3-yl}-acetic acid;
- 25 {5-Methoxy-6-[5-(3-methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-benzofuran-3-yl}-acetic acid;
- {6-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-5-methoxy-benzofuran-3-yl}-acetic acid;
- {6-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-5-methoxy-benzofuran-3-yl}-acetic acid;
- 30 {7-[5-(4-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-indan-4-yloxy}-acetic acid;

{7-[5-(3-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-indan-4-yloxy}-acetic acid;

{7-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-indan-4-yloxy}-acetic acid;

{7-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-indan-4-yloxy}-acetic
5 acid;

{4-[5-(4-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-2-methyl-phenoxy}-acetic acid;

{4-[5-(3-Methoxymethyl-phenyl)-isoxazol-3-ylmethoxy]-2-methyl-phenoxy}-acetic acid;

{4-[5-(4-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-2-methyl-phenoxy}-acetic
acid;

10 {4-[5-(3-Methanesulfonyloxy-phenyl)-isoxazol-3-ylmethoxy]-2-methyl-phenoxy}-acetic
acid; and pharmaceutically acceptable salts thereof.